IN THE CLAIMS

The claims are as follows:

1-172. (Canceled).

173. (Previously Presented) A therapeutic method for treating a cardiovascular or vascular indication characterized by a decreased lumen diameter comprising locally administering to a human identified as being at risk of or afflicted with said cardiovascular or vascular indication, a cytostatic dose of a therapeutic agent, wherein the therapeutic agent is a compound of formula (I):

$$(R^{1})(R^{2})N(CH_{2})_{2}O$$

$$(Z)$$

$$R^{3}$$

$$R^{5}$$

$$R^{6}$$

$$R^{4}$$

$$(I)$$

wherein Z is C=O or a covalent bond; Y is H or $O(C_1-C_4)$ alkyl, R^1 and R^2 are individually (C_1-C_4) alkyl or together with N are a saturated heterocyclic group, R^3 is ethyl or chloroethyl, R^4 is H, R^5 is I, $O(C_1-C_4)$ alkyl or H and R^6 is I, $O(C_1-C_4)$ alkyl or H with the proviso that when R^4 , R^5 , and R^6 are H, R^3 is not ethyl; or a pharmaceutically acceptable salt thereof.

174. (Previously Presented) The method of claim 173 wherein the cytostatic dose is effective to increase the level of TGF-beta so as to decrease lesion formation or development, decrease lipid accumulation, increase plaque stability, maintain or increase vessel lumen diameter, or any combination thereof.

Page 3

Title: PREVENTION AND TREATMENT OF CARDIOVASCULAR PATHOLOGIES WITH TAMOXIFEN ANALOGUES

175. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is

idoxifene, 4-iodotamoxifen, 3-iodotamoxifen, toremifene, or a pharmaceutically acceptable salt

thereof.

176. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is

idoxifene or a pharmaceutically acceptable salt thereof.

177. (Previously Presented) The method of claim 173 wherein the compound of formula (I) is

toremifene or a pharmaceutically acceptable salt thereof.

178. (Canceled).

(Previously Presented) The method of claim 173 wherein the compound of formula (I) is 179.

administered via a sustained release dosage form.

180. (Previously Presented) The method of claim 173 wherein the administration is localized

at the site of vascular trauma.

181. (Previously Presented) The method of claim 173 wherein the compound directly or

indirectly increases the level of active TGF-beta.

182. (Previously Presented) A therapeutic method of increasing the level of TGF-beta in a

diabetic mammal at risk of or afflicted with a cardiovascular or vascular indication characterized

by a decreased lumen vessel diameter, comprising administering to a diabetic mammal at risk of

or afflicted with said cardiovascular or vascular indication an effective amount of a compound of

formula (I):

Page 4 Dkt: 295.009US3

$$(R^1)(R^2)N(CH_2)_2O$$

$$(Z)$$

$$R^3$$

$$R^5$$

$$R^4$$

$$(I)$$

wherein Z is C=O or a covalent bond; Y is H or $O(C_1\text{-}C_4)$ alkyl, R^1 and R^2 are individually $(C_1\text{-}C_4)$ alkyl or together with N are a saturated heterocyclic group, R^3 is ethyl or chloroethyl, R^4 is H or together with R^3 is -CH₂-CH₂- or -S-, R5 is I, OH, $O(C_1\text{-}C_4)$ alkyl or H and R^6 is I, $O(C_1\text{-}C_4)$ alkyl or H with the proviso that when R^4 , R^5 , and R^6 are H, R^3 is not ethyl; or a pharmaceutically acceptable salt thereof.

- 183. (Previously Presented) The method of claim 182 wherein the increase in TGF-beta reduces diabetic retinopathy.
- 184. (Previously Presented) The method of claim 182 wherein the mammal is a human.
- 185. (Previously Presented) The method of claim 184 wherein the diabetic has retinopathy.
- 186. (Previously Presented) The method of claim 182 wherein the compound indirectly or directly increases the level of active TGF-beta in vascular tissue.
- 187. (Previously Presented) The method of claim 182 wherein the compound is a TGF-beta production stimulator.
- 188. (Previously Presented) The method of claim 182 wherein the compound is a TGF-beta activator.

Title: PREVENTION AND TREATMENT OF CARDIOVASCULAR PATHOLOGIES WITH TAMOXIFEN ANALOGUES

189. (Previously Presented) The method of claim 182 wherein the compound increases the production of TGF-beta mRNA.

190. (Previously Presented) The method of claim 182 wherein the compound increases the

cleavage of the latent form of TGF-beta.

191. (Previously Presented) The method of claim 182 wherein the compound increases the

bioavailability of TGF-beta.

192. (Previously Presented) The method of claim 182 wherein the compound is idoxifene or a

pharmaceutically acceptable salt thereof.

193. (Previously Presented) The method of claim 182 wherein the compound is toremifene or

a pharmaceutically acceptable salt thereof.

194. (Previously Presented) The method of claim 182 wherein the compound is droloxifene or

a pharmaceutically acceptable salt thereof.

195. (Canceled)

196. (Previously Presented) The method of claim 173 or 182 wherein the compound forms

cellular DNA adducts at level which is reduced relative to DNA adduct formation by tamoxifen.

197. (Previously Presented) The method of claim 173 or 182 wherein the compound has

estrogenic activity which is reduced relative to the estrogenic activity of tamoxifen.

198. (Previously Presented) The method of claim 173 or 182 wherein the compound does not

form cellular DNA adducts.

Page 6 Dkt: 295.009US3

Title: PREVENTION AND TREATMENT OF CARDIOVASCULAR PATHOLOGIES WITH TAMOXIFEN ANALOGUES

199. (Previously Presented) The method of claim 173 or 182 wherein the compound has no estrogenic activity.

- 200. (Previously Presented) A method of increasing the level of TGF-beta in a human identified as being afflicted with a cardiovascular indication characterized by a decreased lumen vessel diameter, comprising selecting an agent that is structural analog of tamoxifen or a pharmaceutically acceptable salt thereof that directly or indirectly elevates the level of active TGF-beta1 in a human and administering to a human identified as being afflicted with a cardiovascular indication an effective amount of the agent
- 201. (Canceled).
- 202. (Previously Presented) The method of claim 200 wherein the agent is idoxifene or a pharmaceutically acceptable salt thereof.
- 203. (Previously Presented) The method of claim 200 wherein the agent is toremifene or a pharmaceutically acceptable salt thereof.
- 204. (Canceled).
- 205. (Previously Presented) The method of claim 173, 182, or 200 wherein the administration increases the level of latent TGF-beta relative to the level of latent TGF-beta prior to said administration.
- 206. (Previously Presented) The method of claim 173, 182, or 200 wherein the administration increases the level of active TGF-beta relative to the level of active TGF-beta prior to said administration.
- 207-230. (Canceled).

Page 7 Dkt: 295.009US3

231. (Previously Presented) A therapeutic method for treating a condition selected from the group consisting of arteriosclerosis, silent myocardial infarction, vascular insufficiency in the limbs, peripheral neuropathy, and retinopathy, comprising administering to a mammal afflicted with said condition, an effective amount of a compound of formula (I):

$$(R^{1})(R^{2})N(CH_{2})_{2}O$$

$$(Z)$$

$$R^{3}$$

$$R^{5}$$

$$R^{6}$$

$$R^{4}$$

$$(I)$$

wherein Z is C=O or a covalent bond; Y is H or $O(C_1-C_4)$ alkyl, R^1 and R^2 are individually (C_1-C_4) alkyl or together with N are a saturated heterocyclic group, R^3 is ethyl or chloroethyl, R^4 is H, R^5 is I, $O(C_1-C_4)$ alkyl or H and R^6 is I, $O(C_1C_4)$ alkyl or H with the proviso that when R^4 , R^5 , and R^6 are H, R^3 is not ethyl; or a pharmaceutically acceptable salt thereof.

232-233. (Canceled).

234. (Previously Presented) The method of claim 173, 182, or 231 wherein R⁵ or R⁶ is I.